The Use of Multiple "Omic" Platforms to Evaluate the Effects of Endocrine Disrupting Compounds in Small Fish Species

Kimberly J. Ralston-Hooper¹, Gordon J. Getzinger², Meredith E. Turner³, Erik J. Soderblom³, M. Arthur Moseley³, B. Lynn Escalon⁴, Natalia Garcia-Reyero⁵, Carlie A. LaLone⁶, Daniel L. Villeneuve⁶, Gerald T. Ankley⁶, Robert A. Hoke⁷, P. Lee Ferguson²

¹DuPont Visiting Research Scientist at Duke University, Durham NC

Linking molecular changes at multiple levels of biological organization using "omic" methods provides highly complimentary, "data-dense" information for predicting outcomes for organisms exposed to environmental contaminants. However, performing separate "omic" analyses on multiple organisms or tissues increases the variability in responses and the uncertainty of the linkages between these levels of organization. Ideally, performing transcriptomic, proteomic, and metabolomic analyses on the same sample of individual tissues of interest from an organism would reduce this uncertainty and biological variation. On-going research in our laboratory has demonstrated the ability to apply multiple "omic" technologies to analyze individual tissues of small fish species. In the present study, transcriptomic and proteomic analyses were performed on hepatic tissues from individual adult female fathead minnows (FHM; Pimephales promelas) exposed to the aromatase inhibitor fadrozole. Organisms were exposed to 0, 0.04, and 1.0 µg/L fadrozole for 4 days and tissue extracts were analyzed using an LC-MS/MS based, label-free proteomics approach that identified over 1000 proteins. Many of the differentially-expressed proteins in fadrozole-exposed female livers were consistent with published results for fadrozole. Interpretation of the broader proteomic response will be discussed in the context of transcriptomic data collected using mRNA isolated from the same samples used in proteomic analyses. Our data demonstrates the ability to extract high quality RNA and protein samples from small quantities (10 mg) of single tissues for comprehensive 'omic' analyses. Overall, our ability to perform multiple "omic" analyses on individual tissues of small fish species provides greatly improved information for connecting various levels of biological organization and elucidating pathways of toxicity. This information provides a more comprehensive representation of the "systems effects" resulting from chemical exposures and will ideally serve as a basis for advancements in predictive ecotoxicology.

²Department of Civil Engineering and Nicholas School of the Environment, Duke University, Durham NC

³Proteomics Core Facility, Duke University School of Medicine, Durham NC

⁴US Army Engineer Research and Development Center, Vicksburg MS

⁵Institute for Genomics Biocomputing and Bioinformatics, Mississippi State University, Starkville MS

⁶US EPA, National Health and Environmental Effects Research Laboratory, Mid-Continent Ecology Division, Duluth MN

⁷Dupont Haskell Global Centers, Newark DE

STICs Field	Entry
1 – Influence/profile	Not applicable
2 – Clearance tracking no.	Assigned automatically
3 – Principal Investigator /	Kim Ralston-Hooper
Project Officer	Lead EPA Contact – Dan Villeneuve
4- Product title	Copy and paste from abstract
5 - Authors	See abstract
6a- Product type	Presentations and technical summaries
6b-Product subtype	Abstract
6c – Records schedule	Not a senior official
7a – Impact statement	n/a
7b- Product description	Paste in abstract
8 – Bibliographic citation	SETAC North America 33rd Annual Meeting, 11-15 November, Long
	Beach, CA, USA.
9 - Access	Public
10 – Tracking and Planning	2.1.1 2.1.1: Adverse outcome pathway (AOP) discovery and
Task	definition
	definition
10 – Tracking and Planning	(2) AOP descriptions comparing linkages (e.g., causal) between specific
Product	pathway perturbations and reproductive or developmental outcomes in
	multiple species (e.g., rodents, fish, invertebrates) (reports). These will
	provide data that support the development of tools and guidance cross-
	species extrapolation of effects and hazard.
11 – Copyright permission	No .
12 - QA	not applicable
13 – Policy implications	No
14 - Keywords	transcriptomics
	proteomics
	metabolomics
	steroidogenesis

Author	e-mail
Kim Ralston-Hooper	Kimberly.ralston.hooper@duke.edu
Gordon Getzinger	gordon.getzinger@duke.edu
Meredith Turner	Meredith.turner@duke.edu
Erik Soderblom	erik.soderblom@duke.edu
M. Arthur Moseley	Arthur.moseley@duke.edu
B. Lynn Escalon	lynn.escalon@usace.army.mil
Natalia Garcia-Reyero	natalia@icnanotox.org
Robert Hoke	Robert.A.Hoke@USA.dupont.com
P. Lee Ferguson	lee.ferguson@duke.edu